

ANIMAL ALLERGENS, AP HORSE HAIR AND DANDER - equus caballus hair and equus caballus dander injection, solution

ANIMAL ALLERGENS, AP CATTLE HAIR AND DANDER - bos taurus hair and bos taurus dander injection, solution

ANIMAL ALLERGENS, AP DOG HAIR AND DANDER CANIS SPP - canis lupus familiaris hair and canis lupus familiaris dander injection, solution

ANIMAL ALLERGENS, DOG HAIR AND DANDER CANIS SPP. - canis lupus familiaris hair and canis lupus familiaris dander injection, solution

ANIMAL ALLERGENS, FEATHER MIX - gallus gallus feather, anas platyrhynchos feather and anser anser feather injection, solution

ANIMAL ALLERGENS, GUINEA PIG HAIR AND DANDER - cavia porcellus hair and cavia porcellus dander injection, solution

AP HOUSE DUST MIX - house dust injection, solution

DUST, HOUSE MIXTURE - house dust injection, solution

FOOD - ANIMAL PRODUCTS AND POULTRY PRODUCTS, BEEF BOVINE SPP. - beef injection, solution

FOOD - ANIMAL PRODUCTS AND POULTRY PRODUCTS, CHICKEN MEAT GALLUS SP. - poultry injection, solution

FOOD - ANIMAL PRODUCTS AND POULTRY PRODUCTS, EGG, WHITE GALLUS SP. - egg white injection, solution

FOOD - ANIMAL PRODUCTS AND POULTRY PRODUCTS, EGG, YOLK GALLUS SP. - egg yolk injection, solution

FOOD - ANIMAL PRODUCTS AND POULTRY PRODUCTS, PORK SUS SP. - pork injection, solution

FOOD - DAIRY PRODUCTS, CASEIN, COW MILK - casein injection, solution

FOOD - DAIRY PRODUCTS, MILK, WHOLE COW - cow milk injection, solution

FOOD - FISH AND SHELLFISH, CLAM - clam injection, solution

FOOD - FISH AND SHELLFISH, CODFISH GADUS CALLARIAS - codfish injection, solution

FOOD - FISH AND SHELLFISH, CRAB XIPHOSURUS SOWERBYI - crab injection, solution

FOOD - FISH AND SHELLFISH, LOBSTER HOMARUS AMERICANUS - lobster injection, solution

FOOD - FISH AND SHELLFISH, SALMON SALMO SALAR - salmon injection, solution

FOOD - FISH AND SHELLFISH, SHRIMP CRAGO SP. - shrimp injection, solution

FOOD - FISH AND SHELLFISH, TUNA THUNNUS SP. - tuna injection, solution

FOOD - PLANT SOURCE, ALMOND PRUNUS AMYGDALUS - almond injection, solution

FOOD - PLANT SOURCE, APPLE MALUS SP. - apple injection, solution

FOOD - PLANT SOURCE, BANANA MUSA SAPIENTUM - banana injection, solution

FOOD - PLANT SOURCE, BRAZIL NUT BERTHOLLETIA EXCELSA - brazil nut injection, solution

FOOD - PLANT SOURCE, CARROT DAUCUS CAROTA - carrot injection, solution

FOOD - PLANT SOURCE, CASHEW NUT ANACARDIUM OCCIDENTALIE - cashew injection, solution

FOOD - PLANT SOURCE, CELERY APIUM GRAVEOLENS - celery injection, solution

FOOD - PLANT SOURCE, CORN ZEA MAYS - corn injection, solution

FOOD - PLANT SOURCE, HAZELNUT (FILBERT) CORYLUS SPP. - hazelnut injection, solution

FOOD - PLANT SOURCE, MELON, CANTALOUPE CUCUMIS MELO - cantaloupe injection, solution

FOOD - PLANT SOURCE, ORANGE CITRUS SINENSIS - orange injection, solution

FOOD - PLANT SOURCE, PEA, GREEN OR ENGLISH PISUM SATIVUM - pea injection, solution

FOOD - PLANT SOURCE, PEACH PRUNUS PERSICA - peach injection, solution

FOOD - PLANT SOURCE, PEANUT ARACHIS HYPOGAEA - peanut injection, solution

FOOD - PLANT SOURCE, PECAN CARYA ILLINOENSIS - pecan injection, solution

FOOD - PLANT SOURCE, POTATO, WHITE SOLANUM TUBEROSUM - potato injection, solution

FOOD - PLANT SOURCE, RICE, WHOLE GRAIN - rice injection, solution

FOOD - PLANT SOURCE, RYE GRAIN - rye injection, solution

FOOD - PLANT SOURCE, SOYBEAN GLYCINE SOJA - soybean injection, solution

FOOD - PLANT SOURCE, STRAWBERRY FRAGARIA CHILOENSIS - strawberry injection, solution

FOOD - PLANT SOURCE, STRING BEAN MIX - string bean injection, solution

FOOD - PLANT SOURCE, TOMATO NICOTIANA SPP. - tomato injection, solution

FOOD - PLANT SOURCE, WALNUT, BLACK JUGLANS NIGRA - black walnut injection, solution

FOOD - PLANT SOURCE, YEAST, BAKER SACCHAROMYCES CEREVISIAE - yeast injection, solution

FOOD - PLANT SOURCE, YEAST, BREWER SACCHAROMYCES CEREVISIAE - yeast injection, solution

INSECTS (WHOLE BODY) COCKROACH MIX - periplaneta americana and blatella germanica injection, solution

INSECTS (WHOLE BODY), ANT, FIRE SOLENOPSIS INVICTA - solenopsis invicta injection, solution

INSECTS (WHOLE BODY), ANT, FIRE SOLENOPSIS RICHTERI - solenopsis richteri injection, solution

INSECTS (WHOLE BODY), FIRE ANT MIX - solenopsis richteri and solenopsis invicta injection, solution

MOLDS - ALTERNARIA/HORMODENDRUM MIX - alternaria alternata and cladospirium cladosporioides injection, solution

MOLDS - MOLD MIX 10 - *alternaria alternata*, *aspergillus fumigatus*, *emericella nidulans*, *aspergillus niger* var. *niger*, *aspergillus terreus*, *fusarium oxysporum* *vasinfectum*, *dendryphiella vinosa*, *cladosporium cladosporioides*, *mucor racemosus*, *penicillium digitatum*, *penicillium expansum*, *penicillium expansum*, *penicillium chrysogenum* var. *chrysogenum*, *clonostachys rosea* f. *rosea*, *phoma exigua* var. *exigua*, *aureobasidium pullulans* var. *pullulans* and *rhizopus stolonifer* injection, solution

MOLDS - MOLD MIX 4 - *alternaria alternata*, *aspergillus fumigatus*, *emericella nidulans*, *aspergillus niger* var. *niger*, *aspergillus terreus*, *cladosporium cladosporioides*, *penicillium digitatum*, *penicillium expansum*, *penicillium expansum*, *penicillium chrysogenum* var. *chrysogenum* and *clonostachys rosea* f. *rosea* injection, solution

MOLDS - TRICHOPHYTON MIX - *trichophyton tonsurans*, *trichophyton rubrum* and *trichophyton mentagrophytes* injection, solution

MOLDS, PENICILLIUM MIX - *penicillium digitatum*, *penicillium expansum*, *penicillium expansum*, *penicillium chrysogenum* var. *chrysogenum* and *clonostachys rosea* f. *rosea* injection, solution

MOLDS, RUSTS AND SMUTS, ALTERNARIA TENUIS - *alternaria alternata* injection, solution

MOLDS, RUSTS AND SMUTS, ASPERGILLUS FUMIGATUS - *aspergillus fumigatus* injection, solution

MOLDS, RUSTS AND SMUTS, ASPERGILLUS NIGER - *aspergillus niger* var. *niger* injection, solution

MOLDS, RUSTS AND SMUTS, BOTRYTIS CINEREA - *botrytis cinerea* injection, solution

MOLDS, RUSTS AND SMUTS, CANDIDA ALBICANS - *candida albicans* injection, solution

MOLDS, RUSTS AND SMUTS, CEPHALOSPORIUM ACREMONIUM - *acremonium strictum* injection, solution

MOLDS, RUSTS AND SMUTS, CURVULARIA SPICIFERA - *cochliobolus spicifer* injection, solution

MOLDS, RUSTS AND SMUTS, EPICOCUM NIGRUM - *epicoccum nigrum* injection, solution

MOLDS, RUSTS AND SMUTS, EPIDERMOPHYTON FLOCCOSUM - *epidermophyton floccosum* injection, solution

MOLDS, RUSTS AND SMUTS, FUSARIUM VASINJECTUM - *fusarium oxysporum* *vasinfectum* injection, solution

MOLDS, RUSTS AND SMUTS, HELMINTHOSPORIUM INTERSEMINATUM - *dendryphiella vinosa* injection, solution

MOLDS, RUSTS AND SMUTS, HORMODENDRUM CLADOSPORIOIDES - *cladosporium cladosporioides* injection, solution

MOLDS, RUSTS AND SMUTS, MUCOR RACEMOSUS - *mucor racemosus* injection, solution

MOLDS, RUSTS AND SMUTS, PENICILLIUM NOTATUM - *penicillium chrysogenum* var. *chrysogenum* injection, solution

MOLDS, RUSTS AND SMUTS, PHOMA HERBARUM - *phoma exigua* var. *exigua* injection, solution

MOLDS, RUSTS AND SMUTS, PULLULARIA PULLULANS - *aureobasidium pullulans* var. *pullulans* injection, solution

MOLDS, RUSTS AND SMUTS, RHIZOPUS NIGRICANS - *rhizopus stolonifer* injection, solution

MOLDS, RUSTS AND SMUTS, STEMPHYLIUM BOTRYOSUM - *pleospora tarda* injection, solution

POLLENS - GRASSES, BAHIA GRASS PASPALUM NOTATUM - *paspalum notatum* pollen injection, solution

POLLENS - GRASSES, BROME, SMOOTH BROMUS INERMIS - *bromus inermis* pollen injection, solution

POLLENS - GRASSES, CORN, CULTIVATED ZEA MAYS - *zea mays* pollen injection, solution

POLLENS - GRASSES, JOHNSON GRASS SORGHUM HALEPENSE - *sorghum halepense* pollen injection, solution

POLLENS - GRASSES, OATS, COMMON, CULTIVATED AVENA SATIVA - *avena sativa* pollen injection, solution

POLLENS - GRASSES, SOUTHERN GRASS MIX - *poa pratensis* pollen, *dactylis glomerata* pollen, *agrostis gigantea* pollen, *phleum pratense* pollen, *anthoxanthum odoratum* pollen, *sorghum halepense* pollen and *cynodon dactylon* pollen injection, solution

POLLENS - TREES, ACACIA ACACIA LONGIFOLIA - *acacia longifolia* pollen injection, solution

POLLENS - TREES, ALDER, RED ALNUS RUBRA - *alnus rubra* pollen injection, solution

POLLENS - TREES, ASH, WHITE FRAXINUS AMERICANA - *fraxinus americana* pollen injection, solution

POLLENS - TREES, BEECH, AMERICAN FAGUS GRANDIFOLIA - *fagus grandifolia* pollen injection, solution

POLLENS - TREES, BIRCH MIX - *betula papyrifera* pollen, *betula pendula* pollen and *betula nigra* pollen injection, solution

POLLENS - TREES, BOXELDER/MAPLE MIX - *acer negundo* pollen, *acer saccharum* pollen and *acer rubrum* pollen injection, solution

POLLENS - TREES, CEDAR, MOUNTAIN JUNIPERUS ASHEI - *juniperus ashei* pollen injection, solution

POLLENS - TREES, CEDAR, RED JUNIPERUS VIRGINIANA - *juniperus virginiana* pollen injection, solution

POLLENS - TREES, COTTONWOOD, COMMON POPULUS DELTOIDES - *populus deltoides* pollen injection, solution

POLLENS - TREES, CYPRESS, ARIZONA CUPRESSUS ARIZONICA - *cupressus arizonica* pollen injection, solution

POLLENS - TREES, CYPRESS, BALD TAXODIUM DISTICHUM - *taxodium distichum* pollen injection, solution

POLLENS - TREES, ELM, AMERICAN ULMUS AMERICANA - *ulmus americana* pollen injection, solution

POLLENS - TREES, ELM, CHINESE ULMUS PARVIFOLIA - *ulmus parvifolia* pollen injection, solution

POLLENS - TREES, EUCALYPTUS (BLUE GUM) EUCALYPTUS GLOBULUS - *eucalyptus globulus* pollen injection, solution

POLLENS - TREES, GUM, SWEET LIQUIDAMBAR STYRACIFLUA - *liquidambar styraciflua* pollen injection, solution

POLLENS - TREES, HACKBERRY CELTIS OCCIDENTALIS - *celtis occidentalis* pollen injection, solution

POLLENS - TREES, HICKORY, SHAGBARK CARYA OVATA - *carya ovata* pollen injection, solution

POLLENS - TREES, LINDEN (BASSWOOD) TILIA AMERICANA - *tilia americana* pollen injection, solution

POLLENS - TREES, MAPLE, HARD ACER SACCHARUM - acer saccharum pollen injection, solution
POLLENS - TREES, MESQUITE, PROSOPIS JULIFLORA - prosopis juliflora pollen injection, solution
POLLENS - TREES, MULBERRY MIX - morus alba pollen and morus rubra pollen injection, solution
POLLENS - TREES, OAK MIX - quercus rubra pollen, quercus virginiana pollen and quercus alba pollen injection, solution
POLLENS - TREES, OAK, RED QUERCUS RUBRA - quercus rubra pollen injection, solution
POLLENS - TREES, OLIVE OLEA EUROPAEA - olea europaea pollen injection, solution
POLLENS - TREES, PECAN CARYA CARYA ILLINOENSIS - carya illinoensis pollen injection, solution
POLLENS - TREES, PEPPER TREE, CALIFORNIA SCHINUS MOLLE - schinus molle pollen injection, solution
POLLENS - TREES, PINE MIX - pinus contorta pollen and pinus ponderosa pollen injection, solution
POLLENS - TREES, PRIVET LIGUSTRUM VULGARE - ligustrum vulgare pollen injection, solution
POLLENS - TREES, RUSSIAN OLIVE ELAEAGNUS ANGUSTIFOLIA - elaeagnus angustifolia pollen injection, solution
POLLENS - TREES, SYCAMORE, AMERICAN (EASTERN) PLATANUS OCCIDENTALLIS - platanus occidentalis pollen injection, solution
POLLENS - TREES, TREE MIX 11 - fraxinus americana pollen, fagus grandifolia pollen, betula nigra pollen, juglans nigra pollen, populus deltoides pollen, ulmus americana pollen, carya ovata pollen, acer saccharum pollen, quercus rubra pollen, platanus occidentalis pollen and salix nigra pollen injection, solution
POLLENS - TREES, TREE MIX 5 - carya illinoensis pollen, acer saccharum pollen, acer negundo pollen, acer rubrum pollen, quercus rubra pollen, quercus virginiana pollen, quercus alba pollen, platanus occidentalis pollen and salix nigra pollen injection, solution
POLLENS - TREES, TREE MIX 6 - fraxinus americana pollen, fagus grandifolia pollen, betula papyrifera pollen, betula nigra pollen, betula pendula pollen, juglans nigra pollen, populus deltoides pollen and ulmus americana pollen injection, solution
POLLENS - TREES, TREE OF HEAVEN AILANTHUS ALTISSIMA - ailanthus altissima pollen injection, solution
POLLENS - TREES, WALNUT, BLACK JUGLANS NIGRA - juglans nigra pollen injection, solution
POLLENS - TREES, WILLOW, BLACK SALIX NIGRA - salix nigra pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, COCKLEBUR XANTHIUM STRUMARIUM - xanthium strumarium pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, GOLDENROD SOLIDAGO CANADENSIS - solidago canadensis pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, LAMB QUARTERS CHENOPODIUM ALBUM - chenopodium album pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, NETTLE URTICA DIOICA - urtica dioica pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, PIGWEED, ROUGH REDROOT AMARANTHUS RETROFLEXUS - amaranthus retroflexus pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, PLANTAIN, ENGLISH PLANTAGO LANCEOLATA - plantago lanceolata pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, RAGWEED, GIANT AMBROSIA TRIFIDA - ambrosia trifida pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, RAGWEED. WESTERN AMBROSIA PSILOSTACHYA - ambrosia psilostachya pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, RUSSIAN THISTLE SALSOLA KALI - salsola kali pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, SAGEBRUSH, MUGWORT ARTEMISIA VULGARIS - artemisia vulgaris pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, SCALE, WING (SHAD) ATRIPLEX CANESCENS - atriplex canescens pollen injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, SCOTCH BROOM CYTISUS SCOPARIUS - cytisus scoparius injection, solution
POLLENS - WEEDS AND GARDEN PLANTS, SORREL, SHEEP RUMEX ACETOSELLA - rumex acetosella pollen injection, solution
POLLENS - WEEDS, CARELESS WEED AMARANTHUS PALMERI - amaranthus palmeri pollen injection, solution
POLLENS - WEEDS, CARELESS/PIGWEED MIX - amaranthus palmeri pollen and amaranthus retroflexus pollen injection, solution
POLLENS - WEEDS, DOCK/SORREL MIX - rumex crispus pollen and rumex acetosella pollen injection, solution
POLLENS - WEEDS, GIANT, SHORT, WESTERN RAGWEED MIX - ambrosia trifida pollen, ambrosia artemisiifolia pollen and ambrosia psilostachya pollen injection, solution
POLLENS - WEEDS, KOCHIA SCOPARIA - kochia scoparia pollen injection, solution
POLLENS - WEEDS, MARSHELDER/POVERTY MIX - iva axillaris pollen, iva annua var. annua pollen and iva xanthifolia pollen injection, solution

POLLENS - WEEDS, WEED MIX 2630 - xanthium strumarium pollen, chenopodium album pollen, amaranthus retroflexus pollen, rumex crispus pollen and rumex acetosella pollen injection, solution

Hollister-Stier Laboratories LLC

WARNINGS

This product is intended for use only by physicians who are experienced in the use of allergenic extracts, or for use under the guidance of an allergist. Allergenic extracts may potentially elicit a severe life threatening systemic reaction, rarely resulting in death.⁷ Therefore, emergency measures and personnel trained in their use must be available immediately in the event of such a reaction. Patients should be instructed to recognize adverse reaction symptoms and cautioned to contact the physician's office if symptoms occur. Patients on non-selective beta blockers may be more reactive to allergens given for testing or treatment and may be unresponsive to the usual doses of epinephrine used to treat allergic reactions. This product should never be injected intravenously. Refer also to the WARNINGS, PRECAUTIONS, and ADVERSE REACTIONS Sections for further discussion.

DESCRIPTION

Sterile extracts for scratch, prick or puncture testing are supplied in dropper vials containing, in addition to the extract allergens and antigens, 50% (v/v) glycerin as preservative, 0.5% sodium chloride and 0.275% sodium bicarbonate. The strength of these extracts may be expressed in terms of

1. Weight to Volume (w/v)
2. Protein Nitrogen Units/mL (PNU/mL)
3. Allergy Units/mL (AU/mL)
4. Bioequivalent Allergy Units/mL (BAU/mL)
5. Concentrate

1. Weight to volume (w/v).

For regular extracts this describes the extraction ratio, i.e., the amount of crude allergen added to the extracting fluid. A 1:10 extract, therefore, indicates that the solution contains the extracted material from one gram of raw material added to each 10 mL of extracting fluid. The amount and composition of extracted materials will vary with the kind of antigen, the extracting fluid, duration of extraction, pH, temperature, and other variables. APTM (acetone precipitated) extracts, if present, are prepared by reconstituting dry, allergenically active concentrates produced by precipitation process from extracts of raw materials. For those APTM extracts labeled on a weight per volume (w/v) basis, the strength designation indicates the dry weight of finished (acetone) precipitate per volume of reconstituting fluid. For example, 1:50 (w/v) means that each gram of dry precipitate obtained from the original extract is reconstituted in 50 mL of solution.

2. Protein Nitrogen Units per mL (PNU/mL).

One protein nitrogen unit represents 0.00001 mg phosphotungstic acid-precipitable protein nitrogen dissolved in one mL of antigen extract. The PNU content of extracts of the same antigen may vary according to the method of measuring the PNU. Thus, the PNU content of extracts from different manufacturers is not comparable unless the PNU method is known to be the same and reproducible from lot to lot. Also, the amount of protein nitrogen extracted from an antigen is influenced by the same variables as the weight to volume extract. Allergenic materials make up a variable proportion of the total protein of an extract.

3. Allergy Units per mL (AU/mL).

The potency of extracts labeled in Allergy Units per mL (AU/mL) is determined by in vitro comparison to a reference standard established by the Center for Biologics Evaluation and Research (CBER) of the Food and Drug Administration (FDA).

4. Bioequivalent Allergy Units per mL (BAU/mL).

When originally licensed, the Reference Preparations for standardized extracts were arbitrarily assigned 100,000 Allergy Units (AU)/mL. Subsequently, quantitative skin testing by the ID₅₀EAL method 13 was used to determine that some Reference Preparations should be assigned 10,000 AU/mL, and others 100,000 AU/mL. To avoid possible confusion about this change in the method of allergy unit assignment, the nomenclature is being changed for standardized extracts whose allergy units are assigned based on quantitative skin testing, and such products will be labeled in Bioequivalent Allergy Units (BAU)/mL. References labeled 10,000 BAU/mL can be diluted one to a half million fold, and references labeled 100,000 BAU/mL can be diluted one to 5 million fold and produce a sum of erythema diameter of 50 mm when Intradermal testing highly reactive subjects.

5. Concentrate.

Concentrate label terminology applies to allergenic extract mixtures, where the individual allergens being combined vary in strength or the designation of strength.

CLINICAL PHARMACOLOGY

Allergenic extracts for scratch, prick or puncture testing, used according to the DOSAGE AND ADMINISTRATION section, produce erythema or erythema and wheal reactions in patients with significant IgE-mediated sensitivity to the relevant allergen. This allergic inflammatory response, although not completely understood, is thought to begin with reaction of antigen with IgE on the surface of basophils or mast cells, which initiates a series of biochemical events resulting in the production of histamine and other mediators. These, in turn, produce the immediate-type "wheal and flare" skin reaction.

INDICATIONS & USAGE

Certain diagnostics carry labeling which states **Allergenic Extract for Diagnostic Use Only**. Data to support the therapeutic use of products labeled with this statement have not been established. 14

In addition to a carefully taken history, the use of glycerin-containing extracts in scratch, prick or puncture testing is an accepted method in the diagnosis of allergic conditions. 1, 2, 3 Extracts of all allergens do not produce equivalent results in scratch, prick or puncture tests. The intensity of the skin reactions produced will be determined by two factors: the degree of sensitivity of the patient, and the nature of the allergenic extract applied.

Scratch, prick or puncture tests are not as sensitive as the intradermal test, but are safer and cause less discomfort. They may, therefore, be the method of choice when a large number of tests are needed, or when testing the pediatric patient. In some cases, where the relatively insensitive scratch, prick or puncture tests are negative or do not confirm the allergic history, follow-up intradermal tests may be positive. However, **ANTIGENS PRODUCING LARGE 3 to 4+ SCRATCH, PRICK OR PUNCTURE TESTS SHOULD NOT BE TESTED INTRADERMALLY.**

CONTRAINDICATIONS

There are no known absolute contraindications to allergy skin testing. Patients with cardiovascular diseases or pulmonary diseases such as symptomatic asthma, and/or who are receiving cardiovascular drugs such as beta blockers, may be at higher risk for severe adverse reactions. These patients may also be more refractory to the normal anaphylaxis treatment regimen.

WARNINGS

Excessively large local reactions or systemic reactions are more likely to occur if the patient is skin tested shortly after exposure to large amounts of antigen to which s/he is sensitive. Use caution when skin testing patients during a season when pollen is present. Refer to boxed WARNINGS Section.

PRECAUTIONS

1. General

Always have injectable epinephrine and a tourniquet available when tests are being made. (See ADVERSE REACTION section.)

Generally 50 to 60 scratch, prick or puncture tests can be applied safely at one sitting. Patients whose history suggests severe sensitivity should have only 5 to 10 tests applied at a time and these tests applied to the volar surface of one arm. These tests should not all be of the same type of antigen; that is, all grass pollens, all weed pollens, all danders, etc. One or two tests from several classes of antigens should be applied at a time. As soon as a large wheal begins to develop, wipe the antigen from it with a damp cotton sponge. After 10 minutes wipe off all the antigens with a damp cotton sponge, followed by a dry cotton sponge. Be careful not to wipe antigen from a positive reaction onto an adjacent test site.

2 Information for Patients

Patients should be instructed in the recognition of adverse reactions to diagnostic testing. Patients should be made to understand the importance of a 30 minute observation period, and be warned to return to the office promptly if symptoms occur after leaving.

3. Drug Interactions

Patients on non-selective beta blockers may be more reactive to allergens given for testing or treatment and may be unresponsive to the usual doses of epinephrine used to treat allergic reactions.⁶

Certain medications may lessen the skin test wheal and erythema responses elicited by allergens and histamine for varying time periods. Conventional antihistamines should be discontinued at least 5 days before skin testing. Long acting antihistamines should be discontinued for at least 3 weeks prior to skin testing.⁹ Topical steroids should be discontinued at the skin test site for at least 2-3 weeks before skin testing.^{9, 10} Tricyclic antidepressants such as Doxepin should be withheld for at least 7 days before skin testing.¹¹ Topical local anesthetics may suppress the flare responses and should be avoided in skin test sites.¹²

4. Carcinogenesis, mutagenesis, Impairment of Fertility

Long-term studies in animals have not been conducted with allergenic extracts to determine their potential for carcinogenicity, mutagenicity or impairment of fertility.

5. Pregnancy

4,5

Pregnancy Category C. Animal reproduction studies have not been conducted with allergenic extracts. It is also not known whether allergenic extracts can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. Allergenic extracts should be given to a pregnant woman only if clearly needed.

6. Nursing Mothers

There are no current studies on secretion of the allergenic extract components in human milk or effect on the nursing infant. Because many drugs are excreted in human milk, caution should be exercised when allergenic extracts are administered to a nursing woman.

7. Pediatric Use

Wheal sizes in response to allergen skin testing can be smaller in infants than in adults. The skin response to histamine parallels that for allergens; therefore, appropriate positive control skin tests should always be performed.¹

8. Geriatric Use

Skin test wheal size decreases with age. The decrease in allergen-induced skin test reaction parallels that to histamine; therefore, appropriate positive skin test controls should always be performed.¹

ADVERSE REACTIONS

1. Local Reactions

If a severe local reaction occurs during scratch, prick or puncture testing, WIPE OFF test antigen. Large, persistent local reactions or minor exacerbations of the patient's allergic symptoms may be treated by local cold applications and/or the use of oral antihistamines, but they should be considered a warning of possible severe systemic reactions.

2. Systemic Reactions

With careful attention to dosage and administration, such reactions occur infrequently, but it must be remembered that allergenic extracts are highly potent in sensitive individuals and OVERDOSE could result in anaphylactic symptoms. Therefore, it is imperative that physicians administering allergenic extracts understand and be prepared for the treatment of severe reactions.

Adverse reaction frequency data for allergenic extract administration for testing and treatment show that risk is low.^{7, 8}

It cannot be overemphasized that, under certain unpredictable combinations of circumstances, anaphylactic shock is a possibility.

Other possible systemic reaction symptoms include fainting, pallor, bradycardia, hypotension, angioedema, cough, wheezing, conjunctivitis, rhinitis and urticaria. **If a systemic or anaphylactic reaction does occur, WIPE OFF test antigen, apply a tourniquet above the site of injection, if tests are performed on the arms, and inject the 1:1,000 epinephrine-hydrochloride intramuscularly or subcutaneously into the opposite arm. Loosen the tourniquet at least every 10 minutes. Do not obstruct arterial blood flow with the tourniquet.**

EPINEPHRINE:

ADULT DOSAGE: 0.3 to 0.5 mL should be injected. Repeat in 5 to 10 minutes if necessary.

PEDIATRIC DOSAGE: The usual initial dose is 0.01 mg (mL) per kg body weight or 0.3 mg (mL) per square meter of body surface area. Suggested dosage for infants to 2 years of age is 0.05 mL to 0.1 mL; for children 2 to 6 years, 0.15 mL; and children 6 to 12 years, 0.2 mL. Single pediatric doses should not exceed 0.3 mg (mL). Doses may be repeated as frequently as every 20 minutes, depending on the severity of the condition and the response of the patient. After administration of epinephrine, profound shock or vasomotor collapse should be treated with intravenous fluids, and other appropriate drugs. Oxygen should be given by mask. Intravenous antihistamine, theophylline or corticosteroids may be used if necessary after adequate epinephrine and circulatory support have been given. Emergency resuscitation measures and personnel trained in their use should be available immediately in the event of a serious systemic or anaphylactic reaction not responsive to the above measures (Ref. J. Allergy Clin. Immunol. 77 (2): 271-273, 1986). Rarely are all of the above measures necessary; the tourniquet and epinephrine usually produce prompt responses. However, the physician should be prepared in advance for all contingencies. Promptness in beginning emergency treatment measures is of utmost importance.

OVERDOSAGE

See ADVERSE REACTIONS Section.

DOSAGE & ADMINISTRATION

1. General

Parenteral Drug Products should be inspected visually for particulate matter and discoloration prior to administration, whenever solution and container permit.

2. Scratch, Prick or Puncture Testing Methods

There are two general methods of skin testing. (1) The skin is scarified first, and the extract is then applied. (2) A drop of extract is put onto the skin, and a prick or puncture is made through the drop. Avoid touching tip of dropper to skin. Either method is satisfactory, but the second requires that the instrument be cleansed between tests or that separate needles be used.

The extracts for scratch, prick or puncture testing are supplied in dropper vials and should be kept in a rack or box in rows of 10 vials corresponding to the rows of tests to be applied to the skin.

All skin tests should be validated by appropriate positive control tests (e.g., histamine) and negative control tests (e.g., Glycerin, Albumin Saline with Phenol (0.4%), or Buffered Saline with Phenol). The negative control test should be the same material as is used as a diluting fluid in the tested extracts. Diluting fluid is used in the same way as an active test extract.

Test sites should be examined at 15 and 30 minutes. To prevent excessive absorption, wipe off antigens producing large reactions as soon as the wheal appears. Record the size of the reaction. Delayed reactions may rarely occur from tests, so it may be helpful to examine the test sites in 24 hours.

Use of Scarifiers and Spacing. Make scarifications at least 2.5 cm apart. Use more space between pollen tests to prevent smearing into adjacent sites. Hold the scarifier between the thumb and index finger, press the sharp edge of the instrument against the skin and twirl instrument rapidly. The scratch should disrupt only the outer layers of epidermis but should not produce immediate oozing of blood. The amount of pressure needed to produce a satisfactory scratch will vary between patients according to the thickness or fragility of their skin. Experience will indicate the proper amount of pressure to exert in making the scratch. If the scarifier is kept sharp and the scratch made quickly, discomfort to the patient is minimized.

Use of Prick Test Needles. The skin is cleaned and single drops of each extract applied to the properly identified test sites. A small, sterile disposable needle, such as a 1/2-inch 26 gauge needle (with the bevel up), a bifurcated vaccinating needle, or a Prick Lancetter™ is inserted through the drop superficially into the skin, the skin lifted slightly and the needle withdrawn. No bleeding should be produced. After about 1 minute the extract may be wiped away.

3. Most Satisfactory Sites for Testing

Prior to testing, clean the skin area to be tested with ether or alcohol and allow to dry. Use a sterile instrument for each patient. The back or the volar surface of the arms are the most satisfactory sites for testing. Skin of the posterior thighs or abdomen may be used if necessary. Avoid very hairy areas where possible, since the reactions will be smaller and more difficult to interpret. The most satisfactory areas of the back are from the posterior axillary fold to 2.5 cm from the spinal column, and from the top of the scapula to the lower rib margins. The best areas of the arms are the volar surfaces from the axilla to 2.5 or 5 cm above the wrist, skipping the anti-cubital space.

4. Use of Antigen Mixes

The use of complicated mixes of unrelated pollens for testing is not recommended since in the case of a positive reaction, it does not indicate which pollen(s) are responsible, and in the case of a negative reaction, it fails to indicate whether the individual pollens at full concentration would give a positive reaction.

5. Reading Skin Test Reactions

A positive reaction consists of an urticarial wheal with surrounding erythema (resembling somewhat a mosquito bite reaction) larger than the control site. The smallest reaction considered positive is erythema with a central papule at least 5 mm in diameter. In some instances with no reaction at the control site, erythema may be considered an indication of sensitivity. In general, the size of wheal and erythema response correlates directly with the patient's sensitivity to that allergen.

Standardized Products

(a) Mites:

The skin test concentration of 30,000 AU/mL in dropper vials is used for scratch, prick or puncture testing. Puncture tests performed on 12 highly sensitive subjects showed the following:

Species	Mean Sum of Wheal ± Std. Dev. (mm)	Mean Sum of Erythema ± Std. Dev. (mm)
D. farinae	22.4 ± 10.7	82.2 ± 21.7
D. pteronyssinus	24.0 ± 9.9	89.3 ± 24.5

The sum of a skin response is the sum of the longest diameter and the mid-point orthogonal diameter.

(b) Cat Hair and Cat Pelt: The skin test concentration of 10,000 BAU/mL (10-19.9 Fel d 1 Units/mL) in dropper vials is used for prick or puncture testing. Puncture tests performed on 15 highly sensitive subjects showed the following:

Product	Mean Sum of Wheal ± Std. Dev (mm)	Mean Sum of Erythema ± Std. Dev (mm)
Standardized Cat Hair	15.1 ± 3.8	73.3 ± 14.3
Standardized Cat Pelt	13.9 ± 4.3	67.3 ± 13.3

The sum of a skin response is the sum of the longest diameter and the mid-point orthogonal diameter.

(c) Ragweed pollen (Short Ragweed or Giant and Short Ragweed Mixture) Antigen E Assayed: Short Ragweed extract at 1:20 w/v in 50% glycerin containing approximately 100 to 300 units of Antigen E/mL or Giant and Short Ragweed Mix at 1:20 w/v in 50% glycerin containing approximately 50 to 150 units of Antigen E/mL are usually used for scratch, prick or puncture testing. Refer to the following table to determine the skin test sensitivity grade. The corresponding #E (sum of the longest diameter and the mid-point orthogonal diameters of erythema) is also presented.

Grade	Erythema mm	Papule or Wheal mm	Corresponding mm #E
0	<5	<5	<10
±	5-10	5-10	10-20
1+	11-20	5-10	20-40
2+	21-30	5-10	40-60
3+	31-40	10-15 (a)	60-80
4+	>40	>15 (b)	>80

(a) or with pseudopods (b) or with many pseudopods

A positive skin reaction to any allergen must be interpreted in light of the patient's history of symptoms, time of the year, known exposures, and eating habits.

THE SKIN TESTS ARE IN NO WAY A SUBSTITUTE FOR A CAREFUL ALLERGIC HISTORY. THEY SERVE AS ADDITIONAL INFORMATION TO AID IN IDENTIFYING CAUSATIVE ALLERGENS IN PATIENTS WITH ALLERGIC DISORDERS.

6. Geriatric Use

The dose is the same in patients of all age groups. Because the wheal size in response to allergen skin testing decreases with age, appropriate histamine positive control skin tests must be performed.¹

7. Pediatric Use

The dose is the same in patients of all age groups. Wheal size in response to allergen skin testing can be smaller in infants than in adults. Appropriate histamine positive control skin tests must be performed.¹

HOW SUPPLIED

In 5 mL dropper bottles of extract at 1:10 w/v excepting pollens at 1:20, AP™ extracts at 1:50 w/v, except AP™ Dog Hair-Dander at 1:100 w/v, AP™ House Dust at 20,000 PNU/mL, some mixes as Concentrate, and Standardized products at AU/mL (Mite extracts at 30,000 AU/mL) or BAU/mL (Cat Hair and Pelt extracts at 10,000 BAU/mL) value. Strengths are listed on product labels.

STORAGE

The expiration date of the diagnostic extracts is listed on the container label. The extract should be stored at 2° to 8°C and kept at this temperature range during office use.

LIMITED WARRANTY

A number of factors beyond our control could reduce the efficacy of this product or even result in an ill effect following its use. These include storage and handling of the product after it leaves our hands, diagnosis, dosage, method of administration, and biological differences in individual patients. Because of these factors, it is important that this product be stored properly, and that the directions be followed carefully during use. No warranty, express or implied, including any warranty of merchantability or fitness, is made. Representatives of the Company are not authorized to vary the terms or the contents of any printed labeling, including the package insert, for this product except by printed notice from the Company's headquarters. The prescriber and user of this product must accept the terms hereof.

REFERENCES

1. Middleton, Elliott, Jr., C.E. Reed and E.F. Ellis (Eds.) Allergy Principles and Practice. Fourth Edition, Vol. 1. C.V. Mosby. 1993.
2. Sheldon, J.M., R.G. Lovell and K.P. Mathews. A Manual of Clinical Allergy. W.B. Saunders. 1967.
3. Tuft, L. and H.L. Mueller. Allergy in Children. W.B. Saunders. 1970.

4. DuBuske, L.M., C.J. Ling and A.L. Sheffer. Special problems regarding allergen immunotherapy. *Immunol. Allergy Clin. North Am. (USA)*. 12(1): 145-175, 1992.
5. Weinstein, A.M., B.D. Dubin, W.K. Podleski, S.L. Spector and R.S. Farr. Asthma and pregnancy. *JAMA*. 124(11): 1161-1165, 1979.
6. Jacobs, Robert L., Geoffrey W. Rake, Jr., et.al. Potentiated anaphylaxis in patients with drug-induced beta-adrenergic blockade. *J. Allergy Clin. Immunol.* 68(2): 125-127. August 1981.
7. Lockey, Richard F., Linda M. Benedict, Paul C. Turkeltaub, Samuel C. Bukantz. Fatalities from immunotherapy (IT) and skin testing (ST). *J. Allergy Clin. Immunol.* 79(4): 660-677, 1987.
8. Turkeltaub, Paul C., MD, and Peter J. Gergen, MD. The risk of adverse reactions from percutaneous prick-puncture allergen skin testing, venipuncture, and body measurements: Data from the second National Health and Nutrition Examination Survey 1976-80 (NHANES II). *J. Allergy Clin. Immunol.* 84(6): 886-890, Dec. 1989.
9. Pipkorn, Ulf. Pharmacological influence of anti-allergic medication on *In Vivo* allergen testing. *Allergy*. 43: 81-86, 1988.
10. Andersson, M. and U. Pipkorn. Inhibition of the dermal immediate allergic reaction through prolonged treatment with topical glucocorticosteroids. *J. Allergy Clin. Immunol.* 79 (2): 345-349. February 1987.
11. Rao, Kamineni S., et al. Duration of suppressive effect of tricyclic anti-depressants on histamine induced wheal and flare reactions on human skin. *J. Allergy Clin. Immunol.* 82: 752-757, November 1988.
12. Pipkorn, Ulf, and M. Andersson. Topical dermal anesthesia inhibits the flare but not the wheal response to allergen and histamine in the skin prick test. *Clin. Allergy*. 17: 307-311, 1987.
13. Turkeltaub, Paul C., Suresh, C. Rastogi, Harold Baer. Office of Biologics Research and Review skin test method for evaluation of subject sensitivity to standardized allergenic extracts and for assignment of allergy units to reference preparations using the ID₅₀EAL method (Intradermal Dilution for 50 mm Sum of Erythema Determines the Allergy Unit). Methods of the Allergenic Products Branch Office of Biologics Research and Review, FDA, Bethesda, MD 20892. Revised May 9, 1986.
14. Food and Drug Administration. Biological products; Allergenic extracts classified in Category IIIB; Final order; Revocation of licenses. *Federal Register*. 59(220): 59228ff. November 16, 1994.